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### **PATENT COOPERATION TREATY**

|  | n the<br>ERNATIONAL SEA   | ARCHING AUTH        | ORITY   | 1/12                                 |   |  |
|--|---|---------------------|---|--------------------------------------|---|--|
| То   | :   |                     |   | 16/12                                | P () REC'D 0 4 NOV 2004   |  |
|  |   |                     |   | /                                    | WIPO PCT  |  |
|  |   |                     |   |                                      | WIPO PCT  |  |
|  | see form  | PCT/ISA/220         |   | WRIT                                 | TEN OPINION OF THE  |  |
|  |   |                     |   | INTERNATION                          | NAL SEARCHING AUTHORITY   |  |
|  |   |                     |   | (F                                   | PCT Rule 43 <i>bis</i> .1)  |  |
|  |   |                     |   | Data of walling                      |   |  |
|  |   |                     |   | Date of mailing (day/month/year) see | e form PCT/ISA/210 (second sheet)                                   |  |
|  | licant's or agent's file  |                     |   |                                      |   |  |
|  | form PCT/ISA/2  |                     |   | FOR FURTHER A See paragraph 2 below  |   |  |
| Inte   | rnational application   | No                  | International filing date (                           | 1                                    |   |  |
| i  | T/US2004/01717  |                     | International filing date (day/month/year) 28.05.2004 |                                      | Priority date (day/month/year) 04.06.2003                           |  |
| Inte   | mational Patent Clas  | sification (IPC) or | both national classification                          | and IPC                              |   |  |
|  |   |                     | 1,506, A61P25/16                                      | wid ii U                             |   |  |
|  | licant  | ···                 |   |                                      | <del>.</del>  |  |
|  | RCK & CO., INC  | <b>).</b>           |   | •                                    |   |  |
|  |   |                     | · · · · · · · · · · · · · · · · · · ·                 |                                      |   |  |
| 1.   | This opinion co   | ontains indicatio   | ons relating to the folk                              | Owina iteme:                         |   |  |
| "  | _   |                     |   | owing items.                         |   |  |
|  | ☑ Box No. I<br>☑ Box No. II   | Basis of the op     | inion   |                                      |   |  |
|  | Box No. II  | Priority            |   |                                      |   |  |
|  | Box No. IV  | Lack of unity of    |   | ird to novelty, inventive            | e step and industrial applicability                                 |  |
|  | Box No. V   | Reasoned state      | ement under Rule 43 <i>bis</i>                        | .1(a)(i) with regard to r            | novelty, inventive step or industrial                               |  |
|  |   | applicability; cit  | ations and explanations                               | supporting such state                | ment  |  |
|  | ☐ Box No. VI  | Certain docume      |   |                                      |   |  |
|  | ☐ Box No. VII   |                     | In the international app                              |                                      |   |  |
| i _  |   |                     | ations on the internation                             | al application                       | ·   |  |
| 2.   | FURTHER ACTI  | ON                  |   |                                      |   |  |
|  | If a demand for international preliminary examination is made, this opinion will usually be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA"). However, this does not apply where                  |                     |   |                                      | weaver this does not apply where                                    |  |
|  | the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1 bis(b) that written opinions of this International Searching Authority will not be so considered. |                     |   |                                      | onal Searching Authority  |  |
|  | Submit to the IPE   | :A a written reply  | together, where appror                                | riate, with amendmen                 | PEA, the applicant is invited to ts, before the expiration of three |  |
|  | whichever expire  | s later.            | 1 FORK FG 1/13A/220 0[1                               | perore the expiration o              | f 22 months from the priority date,                                 |  |
|  | For further option  | ıs, see Form PC     | T/ISA/220.  |                                      |   |  |
| 3.   | For further details   | s, see notes to F   | orm PCT/ISA/220.                                      |                                      |   |  |
|  |   |                     |   |                                      | ·   |  |
|  | •   |                     |   |                                      |   |  |
|  |   |                     |   |                                      |   |  |
| Name and malling address of the ISA:  Authorized Officer |   |                     |   |                                      |   |  |
|  |   |                     |   |                                      |   |  |

<u>)</u>

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International application No. PCT/US2004/017175

| Box No. I Basis of the opinion  |  |  |  |  |  |
|---|--|--|--|--|--|
| With regard to the language, this opinion has been established on the basis of the international application in the language in which it was field, unless otherwise indicated under this item.   |  |  |  |  |  |
| This opinion has been established on the basis of a translation from the original language into the following language , which is the language of a translation furnished for the purposes of international search (under Rules 12.3 and 23.1(b)).  |  |  |  |  |  |
| With regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed invention, this opinion has been established on the basis of:  |  |  |  |  |  |
| a. type of material:  |  |  |  |  |  |
| ☐ a sequence listing  |  |  |  |  |  |
| ☐ table(s) related to the sequence listing  |  |  |  |  |  |
| b. format of material:  |  |  |  |  |  |
| ☐ in written format   |  |  |  |  |  |
| ☐ in computer readable form   |  |  |  |  |  |
| c. time of filing/furnishing:   |  |  |  |  |  |
| ☐ contained in the international application as filed.  |  |  |  |  |  |
| ☐ filed together with the international application in computer readable form.  |  |  |  |  |  |
| ☐ furnished subsequently to this Authority for the purposes of search.  |  |  |  |  |  |
| 3.  In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished. |  |  |  |  |  |
| . Additional comments:  |  |  |  |  |  |

International application No. PCT/US2004/017175

| Box No. II Priority |   |                   |  |  |  |
|---------------------|---|-------------------|--|--|--|
| 1.                  | 1. ☑ The following document has not been furnished: |                   |  |  |  |
|                     |   | ×                 | copy of the earlier application whose priority has been claimed (Rule 43bis.1 and 66.7(a)).  |  |  |
|                     |   |                   | translation of the earlier application whose priority has been claimed (Rule 43bis.1 and 66.7(b)).   |  |  |
|                     |   | Conser<br>neverth | quently it has not been possible to consider the validity of the priority claim. This opinion has neless been established on the assumption that the relevant date is the claimed priority date.   |  |  |
| 2.                  |   | nas be            | pinion has been established as if no priority had been claimed due to the fact that the priority claim en found invalid (Rules 43 <i>bis.</i> 1 and 64.1). Thus for the purposes of this opinion, the international ate indicated above is considered to be the relevant date. |  |  |
| 3                   | Add   | fitional o        | hservations if necessary   |  |  |

International application No. PCT/US2004/017175

|   | Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability   |   |                                   |  |  |  |  |
|---|--|---|-----------------------------------|--|--|--|--|
| • | he questions whether the claimed invention appears to be novel, to involve an inventive step (to be non bylous), or to be industrially applicable have not been examined in respect of:  |   |                                   |  |  |  |  |
|   | the entire international application,  |   |                                   |  |  |  |  |
| D | claims Nos. 38-46  |   |                                   |  |  |  |  |
| b | because:   |   |                                   |  |  |  |  |
| Σ | the said international application, or the said claims Nos. 38-46 (with respect to industrial applicability) relate to the following subject matter which does not require an international preliminary examination (specify): |   |                                   |  |  |  |  |
|   | see separate sheet   | see separate sheet  |                                   |  |  |  |  |
|   | the description, claims or drawings (indicate particular elements below) or said claims Nos. are so unclear that no meaningful opinion could be formed (specify):  |   |                                   |  |  |  |  |
|   | the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.   |   |                                   |  |  |  |  |
|   | no international search report has been established for the whole application or for said claims Nos.  |   |                                   |  |  |  |  |
|   | the nucleotide and/or amino acid sequence listing does not comply with the standard provided for in Annex C of the Administrative Instructions in that:  |   |                                   |  |  |  |  |
|   | the written form   |   | has not been furnished            |  |  |  |  |
|   |  |   | does not comply with the standard |  |  |  |  |
|   | the computer readable form   |   | has not been furnished            |  |  |  |  |
|   |  |   | does not comply with the standard |  |  |  |  |
|   | the tables related to the nucleo<br>not comply with the technical re   | bles related to the nucleotide and/or amino acid sequence listing, if in computer readable form or<br>emply with the technical requirements provided for in Annex C- <i>bis</i> of the Administrative Instruction |                                   |  |  |  |  |
|   | See separate sheet for further details   |   |                                   |  |  |  |  |

International application No. PCT/US2004/017175

Box No. V Reasoned statement under Rule 43*bis*.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)

Yes: Claims

1-46

No: Claims

Inventive step (IS)

Yes: Claims

Claims

1-46

Industrial applicability (IA)

Yes: Claims

No:

1-37

No: Claims

2. Citations and explanations

see separate sheet

# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY (SEPARATE SHEET)

International application No.

PCT/US2004/017175

Reference is made to the following document:

D1: WO 02/068409 A

#### Re Item III

Claims 38-46 relate to subject-matter considered by this Authority to be covered by the provisions of Rule 67.1(iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject-matter of these claims (Article 34(4)(a)(I) PCT).

#### Re Item V

- 1) The subject-matter of present claims 1-46 is new (Article 33(2) PCT).
- 2) The subject-matter of claims 1-46 does not involve an inventive step (Article 33(3) PCT).

The closest prior art is represented by D1 disclosing NMDA NR2B antagonists (cf. page 1 and claim 1). The structurally closest compounds of D1 are shown in the examples 39 and 40.

The subject-matter of the present claims consists in the selection of a subgroup from the range of compounds described in document D1 (cf. claim 1 of D1). Such a selection can only be regarded as inventive, if the subgroup presents unexpected effects or properties in relation to the rest of the range (i.e. examples 39 and 40). However, no such effects or properties are indicated in the application. Hence, no inventive step is present in the subject-matter of the present claims.